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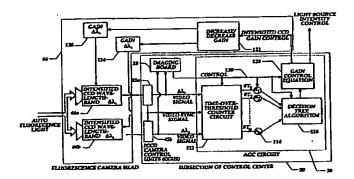
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(54) Title: IMAGING SYSTEM WITH AUTOMATIC GAIN CONTROL FOR REFLECTANCE AND FLUORESCENCE ENDOSCOPY



### (57) Abstract

An imaging system for white light and fluorescence endoscopy that includes an automatic gain control circuit (30) that adjusts the brightness of an image produced based on distribution of pixel intensities in one or more video frames. The magnitude of the image signals produced by a pair of high sensitivity imaging devices such as intensified CCD transducers are compared to a number of reference thresholds. A time-over-threshold counter (112) determines the number of pixels in the image signals having magnitudes greater than or less than the reference thresholds. The distribution of pixel intensities is supplied to a decision tree algorithm (116) that determines whether the gain of the intensified CCD transducers (44a, 44b) used to produce the autofluorescence images or the intensity of the excitation light produced by a light source (36) should be increased or decreased. In addition, a mode switch mechanism is provided to change rapidly from the fluorescence imaging mode to the white light imaging mode or vice versa. This mechanism includes provisions to prevent the accidental application of reflected white illumination light to the image-intensified CCD transducers. Proximity switches (192, 194) monitor the position of a light directing mechanism such as a mirror (186) to allow light to pass to fluorescence camera head (42) or to a color video camera head (46). The light source is not switched to produce white light until it is known that the mirror is in position to direct the reflected light to the color video camera head. Finally, the present invention produces a quantitative display of the relative intensities of the autofluorescence light produced in a pair of spectral bands.

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# IMAGING SYSTEM WITH AUTOMATIC GAIN CONTROL FOR REFLECTANCE AND FLUORESCENCE ENDOSCOPY

# Field of the Invention

The present invention relates to imaging systems for medical endoscopy, in general and to endoscopic imaging systems for fluorescence and reflectance endoscopy, in particular.

# Background of the Invention

One common diagnostic technique used by physicians to detect diseases within a body cavity of a patient is white light optical fiber endoscopy. With this technique, white light is directed into the body cavity via a non-coherent fiber-optic illumination guide of an endoscope. The light illuminates the tissue under examination and the reflected illumination light is gathered and transmitted through a coherent fiber-optic imaging guide of the endoscope. The image formed by the reflected white light at the end of the imaging guide may be viewed directly through the endoscope eyepiece or may be imaged by a color video camera connected to the eyepiece. Images transduced by the camera are then typically transmitted to an image processing/storage device and to a video monitor where they can be viewed by the physician.

To aid physicians performing endoscopy in detecting the presence of cancerous or pre-cancerous tissue, the differences in the autofluorescence (also referred to as native fluorescence) spectrum of normal and abnormal tissue can be exploited. In fluorescence optical fiber endoscopy, a fluorescence excitation light is delivered into the body cavity via the illumination guide of the endoscope. The wavelengths of this light are matched to the absorption spectrum of the naturally

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occurring fluorescing molecules (or fluorophores) present in the tissue (i.e., to blue light). The fluorescence excitation light causes the tissue in the body cavity to fluoresce with a green and red emission spectrum and the resulting light is collected and transmitted through the optical fiber imaging guide of the endoscope. The resulting image is transduced by a camera that filters out any reflected blue light and divides the autofluorescence into two broad (green and red) spectral bands. The image formed by the light in each spectral band is projected onto a separate intensified CCD (ICCD) transducer and the resulting signal is fed into a control center for processing, storage and, finally, for display on a video monitor. The difference in the autofluorescence emission spectrum of normal and abnormal tissue is presented as a difference in color on the video monitor.

Systems for fluorescence fiber endoscopy are fully described in U.S. Patent Nos. 5,507,287, 5,590,660, 5,647,368 and 4,786,813 that are assigned to Xillix Technologies Corp. of Richmond, BC, Canada, the assignee of the present invention, and are sold by Xillix as the Xillix® LIFE-Lung Fluorescence Endoscopy System™ (the "LIFE-Lung System"). Multi-center clinical trials have shown that by using the Xillix LIFE-Lung System as an adjunct to white light endoscopy, the physician's sensitivity in detecting moderate dysplasia, or worse, is 2.71 times greater than the sensitivity of a physician using white light endoscopy alone.

The current LIFE-Lung System has a number of limitations, however. First, the current embodiment of the system requires the physician to manually adjust the gain of the system (i.e., to increase and decrease the camera's sensitivity to the tissue autofluorescence). This is a cumbersome task for the physician to perform, when he/she is simultaneously trying to maneuver the endoscope in the patient. Although automatic gain control circuits for video systems are widely available, they do not provide adequate gain control for the complex scene conditions encountered in imaging autofluorescence with ICCDs. If, for example, the average brightness of an image is increased to an acceptable level, there may be bright spots that can damage the ICCDs. Similarly, if the peak brightness of an image is reduced to prevent localized image saturation, the remainder of the image may become too dark to be recognizable. Furthermore, commonly available average and peak-based automatic gain control circuits do not provide images with a good dynamic range under a variety of viewing conditions, i.e. with an optimized contrast. In endoscopy, these viewing conditions include situations whereby the range of fluorescence light intensities are

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greater than the dynamic range of ICCDs and the image scenes vary from complex structures (i.e. lots of intensity variations) to flat structures (i.e. homogeneous).

A further complication with the use of an automatic gain control circuit arises due to the fact that the gain relationship between the two channels (green and red) of the imaging system must follow a defined function. If the gain of each channel is varied independently, the colors in the resulting video image will not consistently reflect the spectral differences in the autofluorescence of the tissue.

A second limitation of the current LIFE-Lung System becomes evident when a physician wishes to switch between white light (reflectance) and fluorescence imaging modes. With the current system, the physician must switch light sources and cameras manually (i.e., from a white light illumination source to a fluorescence excitation light source and from an RGB color video camera to the fluorescence camera). One technique for addressing this time consuming process is to have all light sources and cameras connected to the endoscope simultaneously and to utilize a mode switching mechanism to switch from one imaging mode to the other. However, some precaution must be taken in the implementation of a switching mechanism since the ICCDs can be damaged if they are subjected to the bright, reflected illumination light. Care must be taken to ensure that the ICCDs are not energized unless the appropriate illumination conditions exist.

A third limitation of the current LIFE-Lung System is that a physician viewing the image displayed by the system has no way of objectively quantifying the extent of abnormality exhibited by the tissue under examination. The effective use of the system is dependent on such subjective factors as the physician's ability to distinguish color and his/her ability to interpret this color information in the context of other image features. A means to objectively quantify the difference in the autofluorescence spectra of normal and abnormal tissue, or even an additional means to subjectively differentiate these tissues based on their difference in autofluorescence spectra could improve the clinical usability of this system. This can be accomplished using computational techniques using the spectral information of the emitted fluorescence and displaying the results on the monitor together with the images.

In summary, the operation of current fluorescence endoscopy systems may be significantly improved by:

a) an automatic gain control circuit that will optimally adjust the brightness of autofluorescence images and that will maintain a defined relationship between the two channels of the imaging system;

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- b) a mechanism that allows rapid switching between white light and fluorescence imaging modes, while preventing the accidental exposure of energized ICCDs to damaging light intensities; and
- c) a means of utilizing the differences in the autofluorescence emission spectra of normal to abnormal tissue to objectively quantify the degree of abnormality of the tissue.

# Summary of the Invention

The present invention is an imaging system for white light and fluorescence endoscopy that includes a particular automatic gain control (AGC) circuit in the fluorescence imaging mode. The AGC circuit adjusts the gain of the imaging system by adjusting the gain of two high sensitivity imaging devices such as image intensified CCD (ICCDs) transducers in a fluorescence camera head and by adjusting the light intensity of the excitation light source. The video signals from a pair channels (the "green" and "red" channel) of a fluorescence camera are supplied to a set of counters. The counters, consisting of counters connected to a clocking oscillator, measure the length of time each video signal has a magnitude that exceeds a reference threshold that is individually set for each counter. Thus, by appropriately arranging the threshold levels, the outputs of the counters can be made to indicate the distribution of video signal amplitudes in one or more video fields. Based upon the outputs of the counters, a decision tree algorithm determines if the gain of the imaging system or the light source intensity should be increased or decreased. A gain control equation determines the appropriate value of light source intensity change and maps the resulting imaging system gain increase or decrease to an individual gain change for each ICCD transducer such that the relative gain between the two channels remains the same.

The present invention also includes a mode switching mechanism that allows for convenient switching between white light and fluorescence endoscopy imaging modes. The implementation of mode switching implies that white light and fluorescence light sources and cameras are connected to the endoscope simultaneously and that the appropriate combination of camera and light source are activated when switching modes. This requires a two-part mode switching mechanism: one switching the cameras and one switching the light sources. The camera mode switching mechanism consists of a light directing mechanism such as a mirror that is movable between a first position, where the image from the endoscope

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is reflected towards an RGB video camera head, and a second position, where the image from the endoscope is allowed to pass to the fluorescence camera head. When a physician uses the mode switch to change from white light imaging to fluorescence imaging or vice versa, a pair of proximity switches provide signals to the system control center, which monitors the position of the mirror, to ensure that the ICCDs are not energized until the appropriate light source has been selected. The light source mode switching mechanism consists of a filter driver that positions blue, fluorescence excitation filters or white light filters in an illumination light path that extends between the light source and an endoscope.

The present invention also provides a means of objectively quantifying the spectral differences between normal and abnormal tissue by using the relative brightness of autofluorescence in the spectral bands being imaged (green and red). A portion of the autofluorescence image is analyzed and the numerical value defined by a particular mathematical function such as the ratio of the image brightnesses of the two wavebands is displayed for the physician.

## Brief Description of the Drawings

The foregoing aspects and many of the attendant advantages of this invention will become more readily appreciated as the same becomes better understood by reference to the following detailed description, when taken in conjunction with the accompanying drawings, wherein:

FIGURE 1 is a block diagram of an imaging system for white light and fluorescence endoscopy according to the present invention;

FIGURE 2 is a block diagram of a light source used in the imaging system for white light and fluorescence endoscopy shown in FIGURE 1;

FIGURE 3 is a block diagram of an automatic gain control circuit in accordance with a first aspect of the present invention;

FIGURE 4 is a block diagram of a number of comparators and time-over threshold counters that are included in the automatic gain control circuit shown in FIGURE 3;

FIGURE 5 is a flowchart illustrating the steps performed by the present invention to change the gain of the imaging system shown in Figure 3 or the intensity of light produced by the light source shown in Figure 2;

FIGURE 6 is a block diagram of an imaging mode switching mechanism located in the combination camera head in accordance with another aspect of the present invention; and

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FIGURE 7 is a pictorial illustration of an autofluorescence image that includes a quantitative indication of the relative intensities of the autofluorescence light present in two spectral bands in accordance with another aspect of the present invention.

## Detailed Description of the Preferred Embodiment

The present invention is an imaging system for white light and fluorescence endoscopy that includes an automatic gain control (AGC) circuit in the fluorescence imaging mode. The AGC circuit controls the image brightness in two ways, a) by adjusting the gain of the two image-intensified CCDs (ICCDs) transducers in a fluorescence camera head, and b) by adjusting the intensity of an excitation light source. The input to the AGC circuit are the two video signals (a green and red channel) produced by the fluorescence camera. The video signals are supplied to a set of counters that determine a total period of time during which the video signal has a magnitude that exceeds reference threshold (set individually for each counter). The outputs of the counters are indicative of the distribution of video signal amplitudes in one or more video fields. Based upon the outputs of the counters, a decision tree algorithm determines if the gain of the imaging system or light source intensity should be increased or decreased. A gain control equation determines the appropriate value of light source intensity change and maps the gain increase or decrease to an individual gain change for each ICCD such that the relative gain between the two channels remains the same.

FIGURE 1 is a block diagram of an imaging system 10 for white light and fluorescence endoscopy according to the present invention. At the heart of the imaging system is a control center 20 that includes a central processing unit 22 that is programmed to control the operation of the system including a combination light source 36 and a combination camera head 42. An internal random access memory (RAM), a hard disk drive and read-only memory (ROM) 24 stores a computer software program that controls the operation of the central processing unit 22. The memory also allows the storage of data such as acquired images, parameters and log files. A number of controls 26 on a front panel of the control center 20, allow an operator to adjust the operation of the imaging system.

The control center 20 also includes an imaging board 28 that receives analog video signals that originate from a number of sources including a fluorescence camera head 44 and an RGB camera head 46 that are enclosed within the combination camera head 42. A video switch, that is part of a digital and video I/O 32, receives and selects the fluorescence or RGB video signals to be supplied as an input to the

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imaging board. The imaging board 28 digitizes the selected video signal, then processes and converts the digitized signals to appropriate signals to be displayed on a video monitor 54.

An automatic gain control circuit 30, included within the control center 20, automatically adjusts the gain of the autofluorescence camera head 44 and the intensity of the fluorescence excitation light from the combination light source 36.

The combination light source 36 provides the white light and fluorescence excitation light. The control center 20 is interfaced with the light source 36 through status and control lines 102, 106, and 108. Broadband white illumination light or fluorescence excitation light (typically at 437nm ±10nm) is supplied from the combination light source 36 to an illumination guide 38 of a fiber-optic endoscope 40. Light from the illumination guide 38 illuminates an internal body cavity of a patient. Reflected white light or autofluorescence light from the tissue under examination is transmitted by an imaging guide of the fiber-optic endoscope 40 and is projected onto the combination camera head 42. The combination camera head 42 also includes a mode switch mechanism 67 that directs the light received from the endoscope 40 to either the RGB video camera head 46 or the fluorescence camera head 44. With the fluorescence imaging mode selected, the fluorescence camera head 44 produces electronic signals that are routed to a dual channel fluorescence camera control unit within the control center 20 (not shown) that converts the electronic signals to standard video signals. The video signals are then routed through the video I/O 32 to the imaging board 28 where they are processed before being displayed on the RGB video monitor 54. Alternatively, if the physician desires to view a reflectance white light image, the position of the mode switch mechanism 67 is selected to project the reflected illumination light onto an RGB video camera head 46. The electronic signals produced by the camera head 46 are supplied to an RGB camera control unit 48 that is external to the control center 20, where they are converted to RGB video signals. The white light RGB video signals are also routed through the video I/O 32 to the imaging board 28 and are processed before being displayed on the RGB video monitor 54. The RGB video camera control unit 48 includes an automatic gain control circuit that also has the capability of adjusting the intensity of the light produced by the combination light source 36 when the system is operating in white light mode. The automatic gain control signals for the white light mode are transmitted to the combination light source on a lead 110.

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A keyboard 52 interfaces with the control center 20 through the digital I/O 32 on the computer motherboard and allows the operator to enter patient data or to change the operating parameters of the imaging system.

In order to display the white light and fluorescence images, the RGB video monitor 54 is connected to the control center 20 through the video I/O 32. A VCR 56 may be connected so that video images can be recorded for later review and analysis. A video printer 58 allows a physician to print hard copies of a video frame. Images may also be recorded by a film recorder 60 or stored on a magneto-optical disk 62.

To allow a user to control the operation of the imaging system, several programmable operator input devices are provided. A footswitch 64 and three operator control switches 65 on the camera head 42 allow the operator to remotely activate various control center 20 functions such as freezing and storing images, selecting different AGC modes, or to control some of the peripheral devices such as the video printer 58, film recorder 60, or magneto optical disk 62.

FIGURE 2 illustrates in further detail the combination light source 36 that is shown in FIGURE 1. The light source includes a metal halide lamp 80 that produces broadband white light with mercury (Hg) peaks. Light produced by the lamp 80 is passed through a number of filters 82. Depending on the imaging mode selected, the light is transmitted through either a broadband white light filter (i.e. triple notch filter to remove the Hg peaks) that eliminates the Hg peaks and shapes the spectrum of the metal halide lamp so that it is similar to that of a Xenon lamp. Alternatively, if fluorescence imaging is selected, light from the lamp is passed through a blue fluorescence excitation light filter that comprises a blue pass band having a center frequency near the mercury peak that occurs at 437 nanometers.

Light passing through the filters 82, also passes through an adjustable intensity control mechanism 84, which controls the intensity of the light delivered to an endoscope. The intensity control 84 is preferably a metal plate with an appropriate shape to block a variable amount of light when it is moved in and out of the light path.

After passing through the intensity control mechanism 84, the light passes through a shutter mechanism 86 that opens to allow the light to enter the illumination guide of the endoscope, if the latter is plugged in.

The operation of the combination light source 36 is controlled by a microprocessor-based light source controller 90. The light source controller 90 controls the operation of a metal halide lamp ballast 92 that provides the operating

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voltage for the metal halide lamp 80. In addition, the light source controller provides control signals to a filter driver 94, that physically moves one of the filters 82 into the light path in accordance with time imaging mode selected.

An intensity control driver 96 receives control signals from the light source controller 90 in order to move the intensity control 84 in and out of the light path, and thereby varies the intensity of the light that reaches the illumination guide of the endoscope. The light source controller 90 also sends control signals to a shutter driver/motor 98 that causes the shutter mechanism 86 to open and close.

In addition to controlling the components that adjust the intensity and wavelength of light that is provided to the illumination guide of the endoscope, the light source controller 90 also interfaces with a number of front panel switches 100 that allow a physician to manually adjust the operation of the light source. Alternatively, the light source controller 90 receives commands to control the light source from an interface to the status and control lines 102 that is coupled to the control center 20 that controls the overall operation of the imaging system as shown in FIGURE 1.

To, change the output of the combination light source 36 from white illumination light to blue excitation light or vice versa, as well as to control the intensity of the light produced, the light source controller 90 also receives control signals from the control center on lead 106 that indicate which of the filters 82 should be placed into the light path in order to create the white light illumination or blue excitation light. The light source controller 90 receives signals from the control center on the status and control lines 108 that indicate whether the intensity of the excitation light produced should be increased or decreased. Finally, the light source controller 90 receives signals from the RGB video camera control unit 48 on the lead 110 that adjusts the intensity of the white illumination light produced.

To eliminate the need for a physician to manually adjust the gain while in the fluorescence imaging mode, the imaging system of the present invention includes a fluorescence mode automatic gain control (AGC) circuit 30 as shown in FIGURE 3. The imaging system can also be operated under manual control as the current Xillix LIFE-Lung Fluorescence Endoscopy System<sup>TM</sup>. The implementation of the fluorescence mode AGC is as follows: As described previously, autofluorescence light produced by the tissue under examination is divided into a pair of spectral bands and projected onto a pair of high sensitivity imaging devices such as a pair of electron bombarded CCD's or image intensified CCD transducers 44a and 44b. The

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transducer 44a receives the light in a wavelength band  $\Delta\lambda_1$ , which is located in the green portion of the visible spectrum, while the transducer 44b receives light in a wavelength band  $\Delta\lambda_2$ , which is located in the red portion of the visible spectrum. The electronic signals produced by the intensified CCD transducers 44a and 44b are supplied to camera control units (CCUs) 45a and 45b within the control center 20, where they are converted into video signals and routed through the video I/O 32 to the imaging board 28 and to the AGC circuit.

The video signals routed to the AGC circuit are applied to a time-overthreshold counter circuit 112. The counter circuit also receives a clock signal which is gated by the horizontal and vertical sync signals from the CCUs. The counter 112 produces a number of outputs #T1, #T2, ... #Tn, each of which contains a value which is proportional to the area in one or more video fields that has an intensity level above an associated predefined threshold intensity value. Each of the output values #T1, ! . #Tn, may be weighted by a function a<sub>1</sub>, ... a<sub>n</sub> 114 before being supplied to a decision tree algorithm 116. The decision tree algorithm 116 determines. if the gain of the imaging system and/or the intensity of the light produced by the combination light source 36 should be increased or decreased. The output of the decision tree algorithm 116 indicates the amount by which the gain should be increased/decreased and this signal is supplied to a gain control equation 120. The gain control calculates the amount by which the light source intensity and/or the gain of the individual intensified CCD transducers 44a and 44b of the imaging system should be adjusted to meet the gain change determined by the decision tree algorithm, while maintaining a predefined gain relationship between the two channels.

If the camera gain is to be increased or decreased, the gain control equation 120 produces a pair of binary numbers whose magnitude will result in a proportional gain change in the two ICCDs. An increase/decrease gain control circuit 122 receives the binary numbers from the gain control equation 120 and converts the binary numbers received into a pair of voltage levels that are supplied to a pair of transducer gain controls 124 and 126. The transducer gain controls 124 and 126 adjust the absolute gain of the intensified CCD transducers 44a and 44b respectively.

FIGURE 4 illustrates in greater detail the time-over-threshold counter 112 described above. The counter 112 operates to produce numeric counts that are indicative of how long a threshold intensity is exceeded in one or more video frames. These numeric counts are proportional to the area in an image with an intensity above

a predefined value. A bank of independently programmable, reference threshold digital-to-analog converters 140 is programmed by the control center 20 to set a series of reference threshold levels against which the video signals from the CCUs are compared. The particular reference threshold levels are selected to represent a percentage of the zero to full scale video signal that is produced by the CCUs and their chosen values are generally dependent on the type of tissue being examined, as will be described below.

The reference thresholds are applied to the inverting inputs of a number of comparators 144. For example, a voltage equal to 45% of the full scale range of the green channel video signal is supplied on a lead 142a to an inverting input of a comparator 144a. Similarly, a voltage equal to 75% of the full scale range is supplied on a lead 142b to an inverting input of a comparator 144b. Another set of reference threshold voltages are applied to a set of comparators that receive the video signal produced by the red channel CCU. In the presently preferred embodiment of the invention, one reference threshold for each channel is set at a desired peak value while the other reference threshold is set at a desired average intensity value:

applied to the noninverting inputs of the comparator circuits 144. When the voltage level of the video signals exceeds the reference thresholds set by the digital-to-analog converters 140, the comparators 144 produce logic high signals. Associated with each of the comparators 144 is a 24-bit counter 146. Each counter has a counter enable pin coupled to the output of its associated comparator such that when the comparator produces the logic high signal, the counter is enabled.

As indicated above, the automatic gain control circuit 30 includes a free running clock 150 having a frequency that is substantially equal to the pixel clock of the CCUs. A sync delay and gating circuit 152 receives the horizontal and vertical synchronization signals produced by the CCUs and only passes the free running clock 150, during the active portions of the video signals. The sync delay and gating circuit 152 also produces a field clock pulse for each field of the video signals received. The pulses are counted by a short counter 154 in order to keep track of the number of field periods associated with the values contained in the time-over-threshold counters.

When the counters 146 are enabled by their corresponding comparator circuits 144, the counters 146 count the number of sync-gated clock pulses that occur during the time when the video signals produced by the red or green channel CCUs

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exceed the reference threshold associated with the comparator that is connected to the counter's enable pin.

The values in the counters 146 are read out through a counter readout control circuit 160 that connects the counters 146 to the imaging system's data bus 130 located on a motherboard within the control center 20. The counter readout control circuit also receives the count held in the short counter 154. The short counter 154 allows the software to be programmed to read out the counters 146 at periodic intervals, such as every ten fields, etc.

Although the presently preferred embodiment of the invention utilizes two reference thresholds for each of the green and red channels, additional threshold counters can be added to the automatic gain control circuit in the manner described above if it is desired to obtain more detailed information on the distribution of the video signal amplitudes.

FIGURE 5 illustrates the steps performed by the decision tree algorithm 116 and the gain control equation 120 shown in FIGURE 3 to adjust the gain of the ICCDs and the light source intensity. FIGURE 5 illustrates the two basic processes used to implement the automatic gain control, namely, i) the setup of the parameters in steps 162 and 164, and ii) the running of the decision tree algorithm and gain control equation in steps 166 to 172.

Beginning with a step 162, the peak and average reference thresholds are set. These values are selected by the operator using the system software. The values selected depend upon the type of image being viewed. In an image that contains many structural features, the thresholds are selected to ensure that all details remain visible. For example, when viewing a body cavity containing detailed structure such as the bronchi, the peak reference threshold may be set at 90% of the full scale value and the average reference threshold set at 50% of the full scale value. Alternatively, if the body cavity being examined is relatively homogeneous, such as the stomach, the reference threshold values may be set such that the average intensity of the image ensures a relatively bright image. For example, the peak reference threshold may be set at 80% of full scale and the average reference threshold set at 60% of full scale. Preprogrammed thresholds selected for commonly viewed tissue samples can be selected or custom values can be entered.

At a step 164, the automatic gain control circuit selects a number of AGC image fill goal values. These values represent the nominal image area for which the video signal amplitude must be greater than or equal to a particular threshold. For

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example, fill goal values may be chosen such that 2% of the image area has video signal amplitudes greater than the peak threshold value and 55% of the image area has video signal amplitudes greater than the average threshold value. The automatic gain control circuit adjusts the gain of the ICCDs and/or the intensity of the light source such that the image intensity distribution calculated from the time-over-threshold counter 112 achieves the best match to the desired image fill goal values. Like the threshold values, the fill goal values are selected by the operator of the system.

Step 166 is the first step in the actual AGC decision tree algorithm. At a step 166, the automatic gain control circuit waits for the last gain change to take effect and then measures the image intensity distribution for specified number fields. At the end of the specified number fields, the values from the counters 146 in the time-over-threshold counter circuit 112 are read and the image areas analyzed.

The image area having video signal amplitudes above the higher, or "peak" threshold, and the image area having video signal amplitudes above the lower, or "average" threshold, are applied to the decision tree at step 168. The decision tree determines whether the gain should be changed so that the intensity distribution will better meet the AGC fill goal values desired. As discussed above, the image area allowed to exceed the peak or the average threshold may be weighted by the functions 114, in order to make the automatic gain control circuit operate more like a peak or average value control circuit as desired for the particular viewing situation.

The amount of gain change determined by the decision tree algorithm 116 is modified by well known process control techniques at a step 170 to optimize transient behavior such as overshoot, settling time, and oscillatory behavior. These techniques include a leaky integrator function, deadband control, control function mapping, proportional control, and rate and range limiting actions on the next applied gain change. These techniques ensure gain changes occur as quickly as possible without creating stability problems.

At a step 172, the gain change for the green or red channel ICCD is determined and if required, the amount of light source intensity change. The gain change is modified by the control techniques and is applied to the gain control equation 120. This equation relates the gain setting of the ICCD in each of the two channels, such that the ratio (first order polynomial) of the gains between the two channels is maintained. The ratio of the gains between the two channels may be selected by the system operator. The operator may adjust the ratio, such that the resulting video image appears more red or more green as desired. In the presently

preferred embodiment of the invention, the relative gain of the ICCD in the red channel to the ICCD in the green channel can be varied over a range of 0.75 to 3. In some applications, the relationship between the gains of the two channels may be a higher order polynomial, e.g.  $g_1 = c + ag_2 + bg_2^2 + ...$  where  $g_1$  is the gain of the red channel,  $g_2$  is the gain of the green channel and a, b, c, are constants.

The situation may occur that the required fluorescence camera gain falls outside of the optimal gain adjustment range of the ICCD in one or both of the channels. If the calculated gain setting of either channel is greater than the maximum optimal setting or smaller than the minimum optimal setting, then the intensity of the excitation light source is increased or decreased by a fixed amount. The intensity of the light produced by the light source is adjusted a sufficient amount to return the camera gain settings to within the optimal working range. A pseudocode listing of the decision tree algorithm 116 and gain control equation 120 is set forth in Appendix A.

The present invention also includes a two part mode switch mechanism (one part in the light source and one part in the combination camera head) that allows for convenient switching between white light and fluorescence endoscopy imaging modes. FIGURE 6 is a schematic block diagram of the mode switching mechanism of the combination camera head. The switching mode mechanism of the light source is shown in FIGURE 2. The preferred embodiment of the mechanism requires the endoscope to be attached to the combination light source 36 and the combination camera head 42 by means of the endoscope connector 180. The combination light source 36 is capable of providing white light (reflectance) illumination and blue light (fluorescence excitation) illumination. The combination camera head 42 is capable of transducing three channel RGB reflectance images and two channel fluorescence images.

Because the light source 36 and camera head 42 are physically separate, the mode switching mechanism is composed of two parts. The two parts of the mechanism are linked through control signals via the imaging system control center 20 and the light source system controller 90. Since the metal halide lamp 80 in the combination light source 36 is capable of providing both the white and blue light, a light source part of the mode switch consists of the filter driver 94 and the white light an blue light filters 82. The filter driver 94 responds to instructions from the light source system controller 90 and positions the appropriate filter in the light path between the lamp and the endoscope illumination guide. The status of the filter driver 94 is also monitored by the light source system controller 90, which then

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communicates with the control center 20 via the interface to the status and control lines 102.

A second part of the mode switching mechanism is located in the combination camera head 42. This part of the mode switching mechanism 67 utilizes a movable light path directing mechanism such as a mirror 186. When in the imaging system is in fluorescence imaging mode, the mirror is moved out of the light path between the endoscope eyepiece and the fluorescence camera head 44. In this way the fluorescence light reaches the dichroic mirror 182 that separates spectrally  $\Delta\lambda_1$  and  $\Delta\lambda_2$  into their respective optical paths. When the imaging system is in the white light imaging mode, the mirror 186 is moved into the light path. In this position, light from the endoscope is directed to a second, fixed mirror 190, where the light path is folded to form a periscope that redirects the light from the endoscope eyepiece to the RGB video camera head 46.

The operation of both parts of the mode switching mechanism is controlled by an operator input on the combination camera head 42. The operator initiates a switch 65 to change the operation of the imaging system. This results in a signal being sent to the control center indicating that a switch of imaging modes has been initiated. A signal is generated by a pair of electrical or optical proximity switches 192, 194 in the combination camera head 42 that sense the position of the movable mirror 186. A second signal is generated by switches 192, 194 and sent to the control center 20 when the movable mirror 186 has reached its new position.

The switches 192, 194 function as a safety mechanism for the ICCDs in the fluorescence camera head. When energized, the ICCDs are susceptible to damage from bright light (e.g. white light reflectance images from the endoscope eyepiece). If the movable mirror 186 is not completely in the fluorescence mode imaging position, the control center 20 reacts by immediately shutting off the power to the ICCDs, thereby protecting them from exposure to possibly damaging illumination.

The control center 20 reacts differently to the switch signals depending on whether the operator is switching from fluorescence imaging mode into the white light imaging mode or from white light imaging mode into the fluorescence imaging mode. In the former case, the control center 20 reacts to the first switch signal by immediately shutting off the power to the ICCDs and stopping the display of all images. When the control center 20 receives the second switch signal, indicating that the movable mirror 186 in the camera head 42 has reached the white light imaging mode position, the control center sends a signal to the light source system

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controller 90 instructing it to move the white light filter into the light path. When the light source mode switch has completed the filter change, the light source system controller 90 generates a light source status signal, which is transmitted to the control center 20. Upon receipt of the light source status signal, the control signal routes the video signal from the RGB video camera control unit 48 to the RGB video monitor 54 and the resulting white light image is displayed.

In the case of switching from white light imaging mode into the fluorescence imaging mode, the control center 20 reacts to the first switch signal from the combination camera head 42 by sending a signal to the light source system controller 90 instructing it to move the blue light filter into the light path. A light source status signal is generated and sent the control center 20 when the light source mode switch has completed the filter change. When the control center also receives the second switch signal from the combination camera head 42, indicating that the movable mirror 186 has reached the fluorescence imaging mode position, the control center energizes the ICCDs in the fluorescence camera head 44 and routes the video signals from the fluorescence camera control units to the RGB video monitor 54. The resulting fluorescence image is displayed on the RGB video monitor 54. If the incorrect light source status signal is received by the control center 20, the ICCDs in the fluorescence camera head will not be energized, even if the second switch signal has been received from the combination camera head 42.

In accordance with another aspect of the present invention, the imaging system of the present invention quantifies the relative brightness of the autofluorescence light produced by the tissue in each of the spectral bands  $\Delta\lambda_1$  and  $\Delta\lambda_2$  in an objective manner. FIGURE 7, shows a monitor display 200 with an image 202 of the tissue under examination. Differences in the autofluorescence spectrum produced by normal and abnormal tissue are shown as areas of different color in the image. For example, abnormal tissue 204 produces proportionally less autofluorescence light in the green portion of the spectrum than normal tissue and is shown as a reddish area in the displayed image.

The relative brightness of the autofluorescence light in the green and red wavebands imaged by the system,  $\Delta\lambda_1$  and  $\Delta\lambda_2$ , can be used as a measure of the difference in the actual fluorescence emission spectra of normal and abnormal tissue. A ratio (or other function relating the  $\Delta\lambda_1$  to the  $\Delta\lambda_2$  waveband) of the brightness of the tissue autofluorescence in the green and red spectral bands is calculated and displayed to the physician. The ratio is calculated for a small area such as a

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region 206 defined in the center of the field of view. Since the color ratio can be recalculated on a frame by frame basis in real time, the color ratio displayed represents the average color ratio of the tissue imaged within the bounds of area 206. Although the area 206 is shown as a particular area located in the center of the field of view, other locations within the field of view and larger or smaller areas could be used.

The ratio calculation is implemented as follows: As described above, the video signals from the fluorescence camera control units are routed to the imaging board 28. The imaging board 28 digitizes the video signals such that the video signal amplitudes correspond proportionally to digital grey level values. The central processing unit 22 within the control center 20 reads the data digitized by the imaging board 28 and sums the grey level values of all the red channel digital data within area 206 and divides that sum by the sum of all the green channel data within area 206. The quotient of these two sums is shown as a dimensionless number 208 on the monitor.

As an alternative to displaying a dimensionless number, other non-visual cues could be used to quantify the relative brightness of the tissue autofluorescence in spectral wavebands  $\Delta\lambda_1$  and  $\Delta\lambda_2$ . For example, a tone having a frequency that is dependent upon the ratio of the brightness of the autofluorescence in each spectral band could be produced. Similarly, the frequency of a blinking light could be made to change in proportion to the changing ratio.

While the preferred embodiment of the invention has been illustrated and described in the preceding description, it will be appreciated that various changes can be made therein without departing from the spirit and scope of the invention. The scope of the invention is therefore to be determined from the following claims and equivalents thereto.

REM PROPERTY OF XILLIX TECHNOLOGIES CORP
REM PROTOTYPE AGC: AUTOCALIBRATION, SET-UP AND TWO COLOR AGC
REM CURTIS FIGLEY, DEC 96, Revised Mar 97

REM BASE ADDRESS OF BOARD HARDWIRED TO D0000 DEF SEG = 53248

REM READ OUT "AGC" LOGO GOSUB READLOGO

REM INITIALIZE DAC'S TO ZERO, SET-UP GAIN AND RANGE GOSUB CLEARDAC

REM MAKESURE ICCD POWER IS OFF
GOSUB HVPSOFF

REM STICK IN SOME STARTING VALUES FOR THE GAIN CONTROL k0! = .72134 / 3: REM PORTION OF GAIN FACTOR OF ~{0.5V/ln(2)} skew! = 1: REM WEIGHTING SHIFT IF deln's ON OPPOSITE SIDES OF '1' REM skew < 1 SHIFTS EMPHASIS TO LARGER ERROR TERM

tlo! = 60: REM LOW THRESHOLD IN PERCENTAGE OF FULLSCALE thi! = 80: REM HIGH THRESHOLD IN PERCENTAGE OF FULLSCALE

flo! = 50: REM NOMINAL PERCENTAGE OF IMAGE ABOVE THRESHOLD ON CHANNEL 5

fhi! = 1.5: REM NOMINAL PERCENTAGE OF IMAGE ABOVE THRESHOLD ON CHANNEL 6

wlo! = 1!: REM FULL AUTO GAIN WEIGHTING FOR CHANNELS (2 & 5)

whi! = 1!: REM FULL AUTO GAIN WEIGHTING FOR CHANNELS (3 & 6)

dband! = .02: REM GAIN DEADBAND IN VOLTS

rg0! = -.014348: rg1! = 1.0494: rg2! = -.00048863#: REM L2PP6 +10% NUMS

steps = 0: REM RED INCREMENTS AROUND NOMINAL

REM CHECK TO SEE IF A PREVIOUS CALIBRATION FILE IS DESIRED

```
300 PRINT "USE PREVIOUS CALIBRATION FILE ('df' for default)?(y/n/q)"
     INPUT bob$: IF (bob$ = "q") THEN 10000
     IF (bob$ = "n") THEN 680
     IF (bob$ = "df") THEN
         name$ = "default.agc"
         GOTO 500
    END IF
     IF (bob$ <> "y") THEN 300
     PRINT "ENTER THE FILENAME TO READ THE DATA FROM"
    INPUT name$
500 OPEN name$ FOR INPUT AS #1
         INPUT #1, npcn2max!, npcn2min!, blklvl2%, whtlvl2%, size2!
         INPUT #1, npcn3max!, npcn3min!, blklvl3%, whtlvl3%, size3!
         INPUT #1, npcn5max!, npcn5min!, blklvl5%, whtlvl5%; size5!
         INPUT #1, npcn6max!, npcn6min!, blklvl6%, whtlvl6%, size6!
     CLOSE #1: GOTO 7500: REM CLOSE INPUT AND SKIP OVER CAL FUNCTIONS
```

# REM INITIALIZE PIXEL COUNTER MAX/MIN's

680 npcn2max! = 0: npcn3max! = 0: npcn5max! = 0: npcn6max! = 0:

z! = 200000: npcn2min! = z!: npcn3min! = z!: npcn5min! = z!

npcn6min! = z!

REM LOOP THROUGH COUNTER READ CYCLES AND PICK OUT MAXIMUM
REM AND MINIMUM COUNTER VALUES...SHOULD CONTAIN MAXIMUM COUNTS
REM IF REFERENCE THRESHOLDS ARE SET BELOW BLACK LEVELS

710 FOR j=0 TO 3: REM GO THROUGH COUNTER SCANNING A FEW TIMES 720 FOR i=0 TO 15: REM VARY # OF FIELDS TO AVOID LATENCY INDUCED ALIASING

A STANDARD TO A STANDARD OF THE STANDARD

730 GOSUB CLEARCNT

REM LET COUNTERS FREE RUN FOR (93 - i) FIELDS
740 fields1! = PEEK(&H0)
IF (fields1! <= (92 - i)) THEN 740

```
REM READ OUT ALL COUNTERS
```

810 GOSUB READCNTS

fields2! = PEEK(&H0)

REM CORRECT FIELDS TO AVERAGE IF FIELDS INCREMENTED DURING READS

Programme of

L HART COLL

fields! = (fields1! + fields2!) / 2

REM CALCULATE TOTAL COUNTS IN EACH CHANNEL

GOSUB MAKECNTS

REM NORMALIZE COUNTER CONTENTS TO NUMBER OF FIELDS TO

REM DETERMINE FULL IMAGE COUNT VALUES

1000 npcn2! = ch2! / fields!: npcn3! = ch3! / fields!

npcn5! = ch5! / fields!: npcn6! = ch6! / fields!

#### REM CHECK FOR UNREALISTIC FULL FILL IMAGE COUNT VALUES

- 1100 flag% = 0; npcn! = 173000; REM npcn! SHOULD BE NOMINAL COUNT VALUE
- 1120 IF (ABS(1 (npcn2! / npcn!)) > .05) THEN flag% = 1
- 1130 IF (ABS(1 (npcn3! / npcn!)) > .05) THEN flag% = 1
- 1150 IF (ABS(1 (npcn5! / npcn!)) > .05) THEN flag% = 1
- 1160 IF (ABS(1 (npcn6! / npcn!)) > .05) THEN flag% = 1
- 1170 IF (flag% > 0) THEN 730: REM GO BACK AND START OVER

#### REM CHECK FOR HIGHER THAN OR LOWER THAN PREVIOUS EXTREMES

1210 IF (npcn2! > npcn2max!) THEN npcn2max! = npcn2!

IF (npcn3! > npcn3max!) THEN npcn3max! = npcn3!

IF (npcn5! > npcn5max!) THEN npcn5max! = npcn5!

IF (npcn6! > npcn6max!) THEN npcn6max! = npcn6!

IF (npcn2! < npcn2min!) THEN npcn2min! = npcn2!

IF (npcn3! < npcn3min!) THEN npcn3min! = npcn3!

IF (npcn5! < npcn5min!) THEN npcn5min! = npcn5!

IF (npcn6! < npcn6min!) THEN npcn6min! = npcn6!

**REM GO AROUND AGAIN** 

1400 NEXT i

1450 NEXT j

REM DETERMINE BLACK LEVEL REFERENCE VALUES BY SLOWLY INCREASING REM REFERENCES UNTIL COUNTS IN EACH CHANNEL FOR DARK IMAGE ARE REM ONE HALF OF FULL COUNTS POSSIBLE ON CHANNEL

REM SET UP FLAGS TO TRIGGER BLACK LEVEL REF VALUE SAVE

1600 flag2 = 0: flag3 = 0: flag5 = 0: flag6 = 0

REM SELECT NEW REFERENCE VOLTAGE

2110 lim% = 260

2120 FOR k% = 130 TO lim%

REM SKIP AHEAD IF ALL THE BLACK LEVELS HAVE BEEN DETERMINED

- 2130 fsum = flag2 + flag3 + flag5 + flag6
- 2140 IF (fsum > 3) THEN k% =  $\lim$ %
- 2150 MSB% = k% \ 16: REM DETERMINE HIGH BYTE lsb% = (k% - (MSB% \* 16)) \* 16: REM DETERMINE HIGH NIBBLE OF LOW BYTE

REM NOW START INCREASING REFERENCE LEVELS ON EACH COUNTER REM UNTIL COUNTS START TO FALL OFF SIGNIFICANTLY

2230 POKE &H31, MSB%: POKE &H30, lsb%: REM REF4 ==> CH6
POKE &H33, MSB%: POKE &H32, lsb%: REM REF3 ==> CH5
POKE &H35, MSB%: POKE &H34, lsb%: REM REF2 ==> CH3
POKE &H37, MSB%: POKE &H36, lsb%: REM REF1 ==> CH2
POKE &H38, 0: REM UPDATE DAC'S

2400 GOSUB CLEARCNT

2420 fields1! = PEEK(&H0): REM WAIT FOR REQUIRED # OF FIELDS

IF (fields1! <= 3) THEN 2420

```
2480 GOSUB READCNTS
```

fields2! = PEEK(&H0)

REM CORRECT FIELDS TO AVERAGE IF IT INCREMENTED DURING CHANNEL READS

fields! = (fields1! + fields2!) / 2

GOSUB MAKECNTS

REM NORMALIZE COUNTER CONTENTS TO NUMBER OF FIELDS

2900 npcn2! = ch2! / fields!: npcn3! = ch3! / fields! npcn5! = ch5! / fields!: npcn6! = ch6! / fields!

REM CHECK TO SEE IF NEW REFERENCE THRESHOLDS HAVE ELIMINATED REM 50% OF POSSIBLE COUNTS ON EACH CHANNEL. IF YES, SET FLAG REM AND ASSIGN CURRENT SETTING AS THIS CHANNELS BLACK LEVEL

3030 IF ((flag2 = 0) AND (npcn2! < (.5 \* npcn2max!))) THEN

flag2 = 1

blklvl2% = k%

END IF

3060 IF ((flag3 = 0) AND (npcn3! < (.5 \* npcn3max!))) THEN

flag3 = 1

blklvl3% = k%

END IF

4020 IF ((flag5 = 0) AND (npcn5! < (.5 \* npcn5max!))) THEN

flag5 = 1

blklvl5% = k%

**END IF** 

4050 IF ((flag6 = 0) AND (npcn6! < (.5 \* npcn6max!))) THEN

flag6 = 1

blklvl6% = k%

END IF

4080 NEXT k%

REM DETERMINE FULLSCALE (WHITE) REFERENCE LEVEL BY RAISING GAINS

REM UNDER USER CONTROL UNTIL SATURATION APPEARS IN EACH COLOR IN REM THE IMAGE. THEN LOCK INTENSIFIER GAINS AND RAISE CHANNEL REM REFERENCEVOLTAGES UNTIL MAJORITY OF COUNTS HAVE BEEN ELIMINATED. REM CORRESPONDING REFERENCE VOLTAGE IS THEN DEFINED AS "FULL SCALE".

#### 4500 GOSUB HVPSON

4520 PRINT "TYPE 'r' WHEN READY, OR 'q' TO QUIT"

INPUT com\$: IF (com\$ = "q") THEN 10000: IF (com\$ <> "r") THEN 4520

REM DETERMINE THE REQUIRED INTENSIFIER SETTING TO
REM PROVIDE SOME SATURATION FOR EACH COLOR CHANNEL

- 4600 PRINT "TYPE 'u' TO RAISE OR 'd' TO DECREASE THE RED GAIN"
  PRINT "TYPE ANY OTHER KEY TO GO ON TO GREEN CHANNEL"
- 4620 msb1% = 100: REM STARTING RED VALUE
- 4630 INPUT dir\$
- 4640 IF ((dir\$ <> "u") AND (dir\$ <> "d")) THEN

  Sat1% = msb1%: REM SAVE CURRENT RED GAIN AS SATURATION GAIN

  POKE &H53, 0: POKE &H58, 0: REM RESET RED GAIN TO ZERO

  GOTO 4800: REM JUMP TO GREEN CHANNEL GAIN SET UP

  END IF
- 4650 IF ((dir\$ = "u") AND (msb1% < 200)) THEN msb1% = msb1% + 4

  IF ((dir\$ = "d") AND (msb1% > 0)) THEN msb1% = msb1% 4

  POKE &H53, msb1%: POKE &H58, 0: GOTO 4630
- 4800 PRINT "TYPE 'u' TO RAISE OR 'd' TO DECREASE THE GREEN GAIN"
  PRINT "TYPE ANY OTHER KEY TO GO ON"
- 4820 msb2% = 100: REM STARTING RED VALUE
- 4830 INPUT dir\$

4840 IF ((dir\$ <> "u") AND (dir\$ <> "d")) THEN

sat2% = msb2%: REM SAVE CURRENT GREEN GAIN AS SATURATION GAIN

POKE &H51, 0: POKE &H58, 0: REM RESET GREEN GAIN TO ZERO

GOTO 5000

END IF

4850 IF ((dir\$ = "u") AND (msb2% < 200)) THEN msb2% = msb2% + 4

IF ((dir\$ = "d") AND (msb2% > 0)) THEN msb2% = msb2% - 4

POKE &H51, msb2%: POKE &H58, 0: GOTO 4830

5000 PRINT "WERE BOTH GAINS SET PROPERLY? (y/n/q)"

INPUT bob\$: IF (bob\$ = "q") THEN 10000

IF (bob\$ = "n") THEN 4500

IF (bob\$ <> "y") THEN 5000

REM IF ALL WAS WELL, RESET SATURATION GAIN VOLTAGES
REM ON RED AND GREEN AND CONTINUE

5060 POKE &H53, sat1%: POKE &H51, sat2%: POKE &H58, 0

REM PAUSE TO LET ICCD HVPS SETTLE

5070 FOR i = 1 TO 20000: NEXT i

REM DETERMINE WHITE LEVEL BY INCREASING THRESHOLDS
REM UNTIL ALMOST ALL COUNTS ARE ELIMINATED

REM SET UP FLAGS TO TRIGGER WHITE LEVEL REF VALUE SAVE

5090 flag2 = 0: flag3 = 0: flag5 = 0: flag6 = 0 max% = 2000

REM SELECT NEW REFERENCE VOLTAGE

5110 FOR k% = 1500 TO max%

REM SKIP AHEAD IF ALL THE WHITE LEVELS HAVE BEEN DETERMINED

5120 fsum = flag2 + flag3 + flag5 + flag6

5130 IF (fsum > 5) THEN k% = max%

MSB% = k% \ 16: REM HIGH BYTE lsb% = (k% - (MSB% \* 16)) \* 16: REM LOW BYTE

POKE &H31, MSB%: POKE &H30, lsb%: REM REF4 ==> CH6

POKE &H33, MSB%: POKE &H32, lsb%: REM REF3 ==> CH5

POKE &H35, MSB%: POKE &H34, lsb%: REM REF2 ==> CH3

POKE &H37, MSB%: POKE &H36, lsb%: REM REF1 ==> CH2

POKE &H38, 0: REM UPDATE DAC'S

#### 5400 GOSUB CLEARCNT

5420 fields1! = PEEK(&H0): REM WAIT FOR REQUIRED # OF FIELDS

IF (fields1! <= 9) THEN 5420

5480 GOSUB READCNTS

fields2! = PEEK(&H0)

REM CORRECT FIELDS TO AVERAGE IF IT INCREMENTED DURING CHANNEL READS

fields! = (fields1! + fields2!) / 2

GOSUB MAKECNTS

REM NORMALIZE COUNTER CONTENTS TO NUMBER OF FIELDS

pcn2! = ch2! / fields!: pcn3! = ch3! / fields! :
pcn5! = ch5! / fields!: pcn6! = ch6! / fields!

REM CHECK TO SEE IF NEW REFERENCE THRESHOLDS HAVE ELIMINATED REM 99% OF POSSIBLE COUNTS ON EACH CHANNEL. IF YES, SET FLAG REM AND ASSIGN CURRENT SETTING AS THIS CHANNELS BLACK LEVEL

6020 IF ((flag2 = 0) AND (pcn2! < (.015 \* npcn2max!))) THEN
flag2 = 1
whtlv12% = k%
END IF

6040 IF ((flag3 = 0) AND (pcn3! < (.015 \* npcn3max!))) THEN flag3 = 1 whtlv13% = k%

```
END IF

6080 IF ((flag5 = 0) AND (pcn5! < (.015 * npcn5max!))) THEN

flag5 = 1

whtlv15% = k%

END IF

6100 IF ((flag6 = 0) AND (pcn6! < (.015 * npcn6max!))) THEN

flag6 = 1

whtlv16% = k%

END IF

6130 NEXT k%
```

REM DETERMINE THE IMAGE SIZE AS A FRACTION OF THE AVAILABLE IMAGE REM AREA BY ADJUSTING THE REFERENCE VOLTAGES TO "JUST" ABOVE BLACK REM AND COUNTING ALL THE PIXELS THAT EXCEED THIS THRESHOLD

REM 'up!' IS THE FRACTIONAL INCREASE (IN TERMS OF THE SIGNAL REM INPUT RANGE) IN THE THRESHOLD ABOVE THE PREVIOUSLY REM DETERMINED BLACK LEVEL (IN MSB)

6200 up! = .05: REM CALCULATE THRESHOLDS CORRESPONDING TO 5% FULLSCALE

lvl2% = blklvl2% + ((whtivl2% - blklvl2%) \* up!)

lvl3% = blklvl3% + ((whtivl3% - blklvl3%) \* up!)

lvl5% = blklvl5% + ((whtivl5% - blklvl5%) \* up!)

lvl6% = blklvl6% + ((whtivl6% - blklvl6%) \* up!)

#### **REM LOAD THRESHOLDS**

POKE &H37, (Ivl2% \ 16): REM CH2 MSB

POKE &H36, ((Ivl2% - ((Ivl2% \ 16) \* 16)) \* 16): REM CH2 LSB

POKE &H35, (Ivl3% \ 16): REM CH3 MSB

POKE &H34, ((Ivl3% - ((Ivl3% \ 16) \* 16)) \* 16): REM CH3 LSB

POKE &H33, (Ivl5% \ 16): REM CH5 MSB

POKE &H32, ((Ivl5% - ((Ivl5% \ 16) \* 16)) \* 16): REM CH5 LSB

POKE &H31, (Ivl6% \ 16): REM CH6 MSB

POKE &H30, ((Ivl6% - ((Ivl6% \ 16) \* 16)) \* 16): REM CH6 LSB

```
POKE &H38, 0: REM UPDATE DAC'S
```

```
6700 cops\% = 0: num\% = 4
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1.00
6710 FOR j\% = 0 TO num%
                                                                                                                                                                                                                                                                                                                                                                                                                                                              医乳糖 医硫酸二磷 化光点量
6730 GOSUB CLEARCNT
6740 fields1! = PEEK(&H0): REM WAIT FOR REQUIRED # OF FIELDS
                                          the contract of the first of th
6780 GOSUB READCNTS
                                                                                                                                                                                                                                                                                                                                                                                                                                                              , 202 (200)
                                                                                                                                                                                                                         A STATE OF THE STA
                                         fields2! = PEEK(\&H0)
                                         REM CORRECT FIELDS TO AVERAGE IF IT INCREMENTED DURING CHANNEL READS
                                         fields! = (fields1! + fields2!) / 2
                                                                                                                                                                                                                                                                                                               The State of the s
                                       GOSUB MAKECNTS
                                                                                                                                                                                                                                                                                                    and the great of the state of t
                                          REM NORMALIZE COUNTER CONTENTS TO NUMBER OF FIELDS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           12. 47 1 - 2350 H
                                          pcnsize2! = ch2! / fields!: pcnsize3! = ch3! / fields!
                                          pensize5! = ch5! / fields!: pensize6!;= ch6! / fields!
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           发展10年中国2014年11月
                                           REM CALCULATE AND CHECK FRACTIONAL SIZE OF IMAGE ON EACH CHANNEL
                                                                                                                                                                               A STATE OF THE STA
   7270 size2! = pcnsize2! / npcn2max!
                                            IF (size2! > 1.1 OR size2! < .2) THEN oops% = 1
   7290 size3! = pcnsize3! / npcn3max!
                                            IF (size3! > 1.1 OR size3! < .2) THEN cops% = 1
   7330 size5! = pcnsize5! / npcn5max!
                                             IF (size5! > 1.1 OR size5! < .2) THEN cops% = 1
   7350 size6! = pcnsize6! / npcn6max!
                                             IF (size6! > 1.1 OR size6! < .2) THEN cops% = 1
   7370 IF (oops% = 0) THEN j\% = num%
    7380 NEXT j%
     7400 IF (oops% <> 0) THEN
                                                                              PRINT "INVALID IMAGE SIZES FOUND"
                                                                                 GOTO 10000
                                              END IF
```

```
REM SHUT DOWN INTENSIFIER GAIN AND POWER
```

· - · · · ·

7430 GOSUB GAINZERO

**GOSUB HVPSOFF** 

**REM PRINT OUT RESULTS** 

7500 PRINT "CHANNEL MAXCNT MINCNT BLKLVL WHTLVL SIZE%"

PRINT " 2 "; npcn2max!; " "; npcn2min!; " "; blklvl2%; " "; whtlvl2%; " "; size2!

PRINT " 3 "; npcn3max!; " "; npcn3min!; " "; blklvl3%; " "; whtlvl3%; " "; size3!

PRINT " 5 "; npcn5max!; " "; npcn5min!; " "; blklvl5%; " "; whtlvl5%; " "; size5!

PRINT " 6 "; npcn6max!; " "; npcn6min!; " "; blklvl6%; " "; whtlvl6%; " "; size6!

#### REM WRITE CURRENT SETTINGS TO FILE

- 7570 PRINT "DO YOU WANT TO SAVE THESE TO A FILE?(y/n/q)"
- 7580 INPUT bob\$: IF (bob\$ = "q") THEN 10000

  IF (bob\$ = "n") THEN 7640

  IF (bob\$ \rightarrow "y") THEN 7570
- 7610 PRINT "ENTER THE FILENAME TO SAVE THE DATA UNDER"
- 7620 INPUT name\$
- 7630 OPEN name\$ FOR OUTPUT AS #1

PRINT #1, npcn2max!, npcn2min!, blklvl2%, whtlvl2%, size2!

PRINT #1, npcn3max!, npcn3min!, blklvl3%, whtlvl3%, size3!

PRINT #1, npcn5max!, npcn5min!, blklvl5%, whtlvl5%, size5!

PRINT #1, npcn6max!, npcn6min!, blklvl6%, whtlvl6%, size6!

CLOSE #1

## REM REBUILD 'pcnsize' VARIABLES

7640 pcnsize2! = size2! \* npcn2max!: pcnsize3! = size3! \* npcn3max!
pcnsize5! = size5! \* npcn5max!: pcnsize6! = size6! \* npcn6max!

REM

REMAGC FEEDBACK DETERMINATION

REM

7680 PRINT " PREPARING TO ENTER IMAGING MODE "

PRINT " PLEASE ESTABLISH THE DESIRED ILLUMINATION"

PRINT " TYPE 'r' WHEN READY, OR 'q' TO QUIT"

INPUT com\$: IF (com\$ = "q") THEN 10000: IF (com\$ <> "r") THEN 7680

7700 PRINT "SELECT GAIN CONTROL METHOD"

PRINT "m -> MANUAL, f -> full (MIXED PEAK/AVERAGE)"

INPUT agcmode\$

IF (agcmode\$ = "m") THEN

PRINT "USE UP/DOWN ARROW KEYS TO CONTROL FINE GAIN"
PRINT "USE PGUP/PGDOWM KEYS TO CONTROL COARSE GAIN"
GOTO 7900

END IF

IF (agcmode\$ = "f") THEN GOTO 7900 GOTO 7700

REM SET UP DEFAULT THRESHOLDS

7900 temp% = CINT((whtlv12% - blklv12%) / 16 \* (tlo! / 100))

POKE &H37, temp%

REM SET REF3 AT ABOUT thi% FS IN MSB

temp% = CINT((whtlv13% - blklv13%) / 16 \* (thi! / 100))

POKE &H35, temp%

REM SET REF5 AT ABOUT tlo% FS IN MSB

temp% = CINT((whtlvi5% - blklvi5%) / 16 \* (tlo! / 100))

POKE &H33, temp%

REM SET REF6 AT ABOUT thi% FS IN MSB

temp% = CINT((whtlv16% - blklv16%) / 16 \* (thi! / 100))

POKE &H31, temp%

POKE &H38, 0: REM UPDATE DAC'S

GOSUB HVPSON

PRINT "ENABLING ICCD HVPS"

now = TIMER

DO WHILE (now = TIMER)

LOOP

REM START CHECKING COUNTERS

delt0! = 1: delt1! = 1: delt2! = 1

CLS

REM SET UP FAULT VARIABLE

warn% = 0: REM CHECKED FOR > 15 FAULTS OUT AND DISABLES ICCD'S

8000 DO WHILE (1 <> 0)

key\$ = INKEY\$

IF ((key\$ = "q") OR (key\$ = "Q")) THEN 9200

IF ((key\$ <> CHR\$(0)) AND (agcmode\$ <> "m")) THEN GOSUB MODVALS: REM TWEAK RUNNING PARAMETERS

REM DELAY AND THEN READ REQUIRED NUMBER OF FIELDS

8070 fields1! = PEEK(&H0)

IF (fields1! <= 3) THEN 8070

**GOSUB CLEARCNT** 

8080 fields1! = PEEK(&H0)

IF (fields1! <= 0) THEN 8080

8180 GOSUB READCNTS

fields2! = PEEK(&H0)

REM CORRECT FIELDS TO AVERAGE IF IT INCREMENTED DURING CHANNEL READS

fields! = (fields1! + fields2!) / 2

**GOSUB MAKECNTS** 

REM NORMALIZE COUNTS TO NUMBER OF FIELDS AND FULL IMAGE COUNTS
REM I.E. 'sx!' IS (AREA ABOVE THRESHOLD/MAXIMUM IMAGE SIZE)

```
s2! = ch2! / fields! / pcnsize2!: s3! = ch3! / fields! / pcnsize3!
     s5! = ch5! / fields! / pcnsize5!: s6! = ch6! / fields! / pcnsize6!
     REM +++++++SIMPLE PSEUDO-PROPORTIONAL AGC WITH DEADBAND+++++++++++++
     REM ++ERROR TERM BASED UPON WEIGHTED AVERAGE OF INDIVIDUAL ERRORS++
    REM CAN'T HAVE MORE THAN ALL THE IMAGE
8700 IF (s2! > 1) THEN s2! = 1: IF (s3! > 1) THEN s3! = 1
     IF (s5! > 1) THEN s5! = 1: IF (s6! > 1) THEN s6! = 1
    REM RELATIVE AMOUNT OF IMAGE IN RELATION TO DESIRED LEVEL
    del2! = (s2! / (flo! / 100))
     del3! = (s3! / (fhi! / 100))
     del5! = (s5! / (flo! / 100))
     del6! = (s6! / (fhi! / 100))
     REM SKIP PAST AUTOMATIC GAIN CHANGE DETERMINATION IF IN MANUAL MODE
     IF (agcmode$ = "m") THEN
         GOSUB MANUALdg
         GOTO 8850
     END IF
     REM CHECK FOR SEVERE OVERSATURATION CONDITIONS
     IF ((del2! > 6) OR (del6! > 6)) THEN
         warn% = warn% + 1: REM CHARGE WARNING VALUE
     ELSE
         warn% = warn% - 1: REM DISCHARGE WARNING VALUE
         IF (warn% < 0) THEN warn% = 0
     END IF
     REM IF SATURATED TO MANY TIMES IN A ROW, CRASH AND BURN
     IF (warn% > 25) THEN
         GOSUB HVPSOFF
         FOR i = 58 TO 74 STEP 4
         SOUND (EXP(i / 10)), 1
```

```
NEXT i
    PRINT "AGC UNABLE TO PREVENT SERIOUS SATURATION"
    PRINT "PLEASE VERIFY SYSTEM SETTINGS AND CONFIGURATION"
    GOTO 9200
END IF
IF (warn% > 16) THEN SOUND 3600, 1
REM ARBITRATE GAIN CONTROL INPUTS BASED UPON
REM RELATIVE RED AND GREEN IMAGE FILLS
IF ((del5! >= del2!) AND (del6! >= del3!)) THEN
    REM GREEN IMAGE CONTAINS HIGHEST RELATIVE IMAGE FILLS THEREFORE
    REM GENERATE A WEIGHTED ERROR VALUE ONLY FROM GREEN VALUES
    IF (SGN(1 - del5!) = SGN(1 - del6!)) THEN
         delt3! = ((wlo! * del5!) + (whi! * del6!))
         delt3! = delt3! / (wlo! + whi!)
         MODE$ = "5AND6"
    END IF
    IF ((del5! < 1) AND (del6! > 1)) THEN
         REM SLIGHTLY SHIFT WEIGHTING TOWARDS CHANNEL 5
         delt3! = ((skew! * wlo! * del5!) + (whi! / skew! * del6!))
         delt3! = delt3! / (skew! * wlo! + whi! / skew!)
         MODE$ = "50VER6"
    END IF
    IF ((del6! < 1) AND (del5! > 1)) THEN
         REM SLIGHTLY SHIFT WEIGHTING TOWARDS CHANNEL 6
         delt3! = ((wlo! / skew! * del5!) + (skew! * whi! * del6!))
         delt3! = delt3! / (wlo! / skew! + skew! * whi!)
         MODE$ = "60VER5"
    END IF
ELSEIF ((del2! >= del5!) AND (del3! >= del6!)) THEN
    REM RED IMAGE CONTAINS HIGHEST RELATIVE IMAGE FILLS THEREFORE
    REM GENERATE A WEIGHTED ERROR VALUE ONLY FROM RED VALUES
    IF (SGN(1 - del2!) = SGN(1 - del3!)) THEN
         delt3! = ((wlo! * del2!) + (whi! * del3!))
```

```
delt3! = delt3! / (wlo! + whi!)
                      MODES = "2AND3"
           END IF
           IF ((del2! < 1) AND (del3! > 1)) THEN
                      REM SLIGHTLY SHIFT WEIGHTING TOWARDS CHANNEL 2
                       delt3! = ((skew! * wio! * del2!) + (whi! / skew! * del3!))
                      delt3! = delt3! / (skew! * wlo! + whi! / skew!)
                      MODE$ = "20VER3"
           END IF
           IF ((del3! < 1) AND (del2! > 1)) THEN
                       REM SLIGHTLY SHIFT WEIGHTING TOWARDS CHANNEL 3
                       delt3! = ((wlo! / skew! * del2!) + (skew! * whi! * del3!))
                       delt3! = delt3! / (wlo! / skew! + skew! * whi!)
                      MODE$ = "3OVER2"
                                                                  END IF
REM IF NEITHER RED OR GREEN IS CLEARLY DOMINANT, MIX RESULTS
REM TO GENERATE AN ERROR TERM
                     the state of the contract of the state of th
ELSEIF ((del5! >= del2!) AND (del3! >= del6!)) THEN
           REM GREEN MID AND RED PEAK
           IF (SGN(1 - del5!) = SGN(1 - del3!)) THEN
                       delt3! = ((wlo! * del5!) + (whi! * del3!))
                       delt3! = delt3! / (wlo! + whi!)
                       MODE$ = "5AND3 "
           END IF
            IF ((del5! < 1) AND (del3! > 1)) THEN
                       REM SLIGHTLY SHIFT WEIGHTING TOWARDS CHANNEL 5
                        delt3! = ((skew! * wlo! * del5!) + (whi! / skew! * del3!))
                        delt3! = delt3! / (skew! * wlo! + whi! / skew!)
                        MODES = "50VER3"
             END IF
             IF ((del3! <= 1) AND (del5! > 1)) THEN
                        REM SLIGHTLY SHIFT WEIGHTING TOWARDS CHANNEL 3
                         delt3! = ((wlo! / skew! * del5!) + (skew! * whi! * del3!))
```

```
delt3! = delt3! / (wlo! / skew! + skew! * whi!)
       MODES = "30VER5"
   END IF
ELSEIF ((del2! >= del5!) AND (del6! >= del3!)) THEN
   REM RED MID AND GREEN PEAK
   IF (SGN(1 - del2!) = SGN(1 - del6!)) THEN
        delt3! = ((wlo! * del2!) + (whi! * del6!))
       delt3! = delt3! / (wlo! + whi!)
       MODE$ = "2AND6"
   END IF
    IF ((del2! < 1) AND (del6! > 1)) THEN
        REM SLIGHTLY SHIFT WEIGHTING TOWARDS CHANNEL 2
       delt3! = ((skew! * wlo! * del2!) + (whi! / skew! * del6!))
        delt3! = delt3! / (skew! * wlo! + whi! / skew!)
        MODE$ = "2OVER6"
   END IF
    IF ((del6! < 1) AND (del2! > 1)) THEN
        REM SLIGHTLY SHIFT WEIGHTING TOWARDS CHANNEL 6
        delt3! = ((wlo! / skew! * del2!) + (skew! * whi! * del6!))
        delt3! = delt3! / (wlo! / skew! + skew! * whi!)
        MODE$ = "6OVER2"
   END IF
ELSE
    REM IF SITUATION GETS TO HERE, MUST HAVE MISSING TEST CASE
    REM RUN IN SMALL CIRCLES AND PANIC
    BEEP
    LOCATE 20, 18
    PRINT "del2="; del2!, " del3="; del3!, " del5="; del5!, " del6="; del6!
    GOTO 10000
END IF
REM FOUR SAMPLE INTEGRATOR WITH GEOMETRIC DECAY '1/{(t-t0)^2}'
REM BUILD delt! FROM RECENT GAIN CHANGE HISTORY OF deltn!'s
```

```
8720 delt! = (delt3! + delt2! / 3 + delt1! / 9 + delt0! / 27)
     delt! = delt! / (1 + 1 / 3 + 1 / 9 + 1 / 27): REM PRESERVE SCALE
     REM SHIFT deltn!'s ONE PERIOD OLDER
     delt0! = delt1!: delt1! = delt2!: delt2! = delt3!
     REM PROTECT 'LOG' FROM NEAR ZERO OR LARGE 'delt!' VALUES
     IF (delt! < .000001) THEN delt! = .000001: REM LIMITS MAXIMUM DOWN STEP.
     IF (delt! > 1000000) THEN delt! = 1000000: REM LIMITS MAXIMUM UP STEP
     REM LOG RESPONSE TO COMPLEMENT (e^V) BEHAVIOUR
     REM NOTE THAT LARGE delt! CAUSES NEGATIVE dg!
     dg! = -1 * LOG(delt!) * k0!
                                                                           ed© kiri
     IF (dg! > 2) THEN dg! = 2: REM OVERRIDE MAXIMUM UP-STEP
     REM NOW APPLY GAIN CHANGE, DEADBAND AND FAILURE LIMITS
8800 IF (ABS(dg!) <= dband!) THEN dg! = 0: REM APPLY DEADBAND
     REM APPLY GAIN CHANGE
     REM -> ASSUMES RED AND GREEN HAVE SIMILAR d/dV{Krel} CHARACTERISTICS
8850 g! = g! + dg!
     IF (g! < .01) THEN g! = .01: REM BOTTOM GAIN LIMIT
     IF (g! > 9.3) THEN g! = 9.3: REM TOP GAIN LIMIT
     gain% = g! * 4096 / 10: REM GREEN GAIN SETTING IN PARTS PER 4096
     REM SEND OUT NEW INTENSIFIER GAIN REQUESTS
8900 grnmsb% = gain% \ 16
     grnlsb\% = (gain\% - (grnmsb\% * 16)) * 16
     r! = rg0! + g! * rg1! + (g! ^2) * rg2!: REM RED GAIN IN VOLTS
     rgain% = CINT((r! * 4096) / 10): REM RED GAIN SETTING IN PARTS PER 4096
     redmsb\% = rgain\% \setminus 16
     redlsb% = (rgain% - (redmsb% * 16)) * 16
     REM SEND OUT NEW CONTROL VOLTAGES
```

9000 POKE &H53, redmsb%: POKE &H52, redlsb%

```
POKE &H51, grnmsb%: POKE &H50, grnlsb%
                         POKE &H58, 0
                         LOCATE 16, 27
                         format3$ = "&##.##&##.##&&"
                                                                                                                                                                                                        Same of the state 
                         COLOR 14
                         LOCATE 19, 20
                                                                                ニー・ディー・エン・コング しない さいし カードはんごうんど 人士
                         COLOR 4
                         format4$ = "&##.##&##.##&##.##"
                         PRINT USING format4$, "del2="; del2!; " del3="; del3!; " del5="; del5!; " del6="; del6!
9100 LOOP
                                                                                                                                                                                            9200 CLS
                         COLOR 7
                         PRINT "SETTING GAINS TO ZERO AND DISABLING ICCD HVPS"
                         GOSUB GAINZERO
                          GOSUB HVPSOFF Death And the state of the sta
9750 PRINT "DO YOU WANT TO QUIT (y/n)?"
                                                                                                                                                                                                                                      INPUT com$
                         IF (com$ = "y") THEN GOTO 10001
                          IF (com\$ = "n") THEN
                                             PRINT "DO YOU WANT TO RESET GAIN TO LAST VALUE (y/n)?"
                                              INPUT com$
                                                                                                                                                                                                                           . .
                                              IF (com\$ = "y") THEN
                                                                  POKE &H53, redmsb%: POKE &H52, redlsb%
                                                                  POKE &H51, grnmsb%: POKE &H50, grnlsb%
                                                                  POKE &H58, 0
                                              END IF
                                              GOTO 7700
                          END IF
                           GOTO 9750
    10000 COLOR 7
```

```
PRINT "SETTING GAINS TO ZERO"
    GOSUB GAINZERO
    PRINT "BYE BYE"
10001 END
READLOGO:
    as = CHRS(PEEK(\&HD)): bs = CHRS(PEEK(\&HE)): cs = CHRS(PEEK(\&HF))
    logo\$ = (a\$ + b\$ + c\$)
    IF (logo$ <> "AGC") THEN
       PRINT " AGC BOARD LOGO NOT FOUND"
       PRINT " PLEASE CHECK EQUIPMENT SET UP"
       PRINT "
                     EXITING"
       GOTO 10000
    END IF
    PRINT " ++++++"
    PRINT " + "; logo$; " +"
    PRINT " ++++++"
    RETURN
                  · 1
CLEARDAC:
    POKE &H21, 0: POKE &H28, 0: REM SET GAIN AND RANGE OF LOW DAC'S
    POKE &H41, 0: POKE &H48, 0: REM SET GAIN AND RANGE OF HIGH DAC'S
    REM DAC MSB DAC LSB
    REM ____
    REM | | |
    REM \ / \ /
    REM \/ \/
    POKE &H37, 0: POKE &H36, 0: REM REF2 ==> CH2\
                         >-RED CHANNELS
REM
    POKE &H35, 0: POKE &H34, 0: REM REF3 ==> CH3/
    POKE &H33, 0: POKE &H32, 0: REM REF5 ==> CH5\
```

```
>-GREEN CHANNELS
REM
     POKE &H31, 0: POKE &H30, 0: REM REF6 ==> CH6/
     POKE &H51, 0: POKE &H50, 0: REM GREEN GAIN CONTROL
     POKE &H53, 0: POKE &H52, 0: REM RED GAIN CONTROL
     POKE &H38, 0: POKE &H58, 0: REM UPDATE DAC'S
                    REM REF1 ==> CH1 **NO LONGER USED**
     REM REF4 ==> CH4 **NO LONGER USED**
     RETURN
CLEARCNT: REM COUNTERS CLEARED BY READ TO HIGHEST LOGO CHARACTERS
     gill% = PEEK(&H1F)
     RETURN
READCNTS:
     ch2high! = PEEK(&HC): ch2mid! = PEEK(&HB): ch2low! = PEEK(&HA)
     ch3high! = PEEK(\&H9): ch3mid! = PEEK(\&H8): ch3low! = PEEK(\&H7)
     ch5high! = PEEK(&H6): ch5mid! = PEEK(&H5): ch5low! = PEEK(&H4)
     ch6high! = PEEK(&H3): ch6mid! = PEEK(&H2): ch6low! = PEEK(&H1)
     RETURN
MAKECNTS:
     ch2! = (((ch2high! * 256) + ch2mid!) * 256) + ch2low!)
     ch3! = ((((ch3high! * 256) + ch3mid!) * 256) + ch3low!)
     ch5! = (((ch5high! * 256) + ch5mid!) * 256) + ch5low!)
     ch6! = ((((ch6high! * 256) + ch6mid!) * 256) + ch6low!)
     RETURN
HVPSON: REM ANY WRITE TO ODD ADDRESS IN RANGE OF &H60 TO &H7F SETS TO ON
```

POKE &H61, 0: REM ENERGIZE ICCD HVPS

RETURN

```
HVPSOFF: REM ANY WRITE TO EVEN ADDRESS IN RANGE OF &H60 TO &H7F SETS TO OFF
     POKE &H60, 0: REM DE-ENERGIZE ICCD HVPS
     RETURN
GAINZERO:
     POKE &H53, 0: POKE &H52, 0
     POKE &H51, 0: POKE &H50, 0
     POKE &H58, 0
     RETURN
MANUALdg:
     dg! = 0: REM DEFAULT GAIN CHANGE
     REM CHECK FOR UP ARROW
     IF (\text{key}) = (\text{CHR}(0) + \text{CHR}(72)) THEN
          dg! = .05
          LOCATE 12, 2
          PRINT "UP "
     END IF
     REM CHECK FOR DOWN ARROW
     IF (\text{key}) = (\text{CHR}(0) + \text{CHR}(80)) THEN
          dg! = -.05
          LOCATE 12, 2
          PRINT "DN "
     END IF
     REM CHECK FOR PGUP
     IF (key$ = (CHR$(0) + CHR$(73))) THEN
          dg! = .5
          LOCATE 12, 2
          PRINT "UPFAST"
     END IF
     REM CHECK FOR PGDOWN
      IF (\text{key}\$ = (\text{CHR}\$(0) + \text{CHR}\$(81))) THEN
          dg! = -.5
```

LOCATE 12, 2

```
PRINT "DNFAST"

END IF

REM OTHERWISE HOLD GAIN AT CURRENT SETTING

IF (dg! = 0) THEN
```

LOCATE 12, 2

PRINT "HOLD "

**END IF** 

RETURN

## MODVALS:

key1S = keyS

REM STRIP key\$ TO SINGLE CHARACTER

key\$ = LEFT\$(key\$, 1)

IF (key\$ > key1\$) THEN BEEP

## REM CHECK FOR WHICH VALUE TO CHANGE

IF (key\$ = "z") THEN

tlo! = tlo! - 1

IF (tlo! < 30) THEN tlo! = 30

REM SET REF2 AT ABOUT 110% FS IN MSB

temp% = CINT((whtlvl2% - blklvl2%) / 16 \* (tlo! / 100))

POKE &H37, temp%

REM SET REF5 AT ABOUT 110% FS IN MSB

temp% = CINT((whtlvl5% - blklvl5%) / 16 \* (tlo! / 100))

POKE &H33, temp%

POKE &H38, 0: REM UPDATE DAC'S

**END IF** 

IF (key\$ = "Z") THEN

tlo! = tlo! + 1

IF (tlo! > 80) THEN tlo! = 80

REM SET REF2 AT ABOUT 110% FS IN MSB

temp% = CINT((whtlvl2% - blklvl2%) / 16 \* (tlo! / 100))

```
POKE &H37, temp%
    REM SET REF5 AT ABOUT 110% FS IN MSB
    temp\% = CINT((whtlv15\% - blklv15\%) / 16 * (tlo! / 100))
    POKE &H33, temp%
    POKE &H38, 0: REM UPDATE DAC'S
END IF
IF (key$ = "x") THEN
    thi! = thi! - 1
    IF (thi! < 50) THEN thi! = 50
    REM SET REF3 AT ABOUT thi% FS IN MSB
    temp% = CINT((whtlv13% - blklv13%) / 16 * (thi! / 100))
    POKE &H35, temp%
    REM SET REF6 AT ABOUT thi% FS IN MSB
    temp% = CINT((whtlvl6% - blklvl6%) / 16 * (thi! / 100))
    POKE &H31, temp%
    POKE &H38, 0: REM UPDATE DAC'S
                                                    7.1
END IF
IF (key$ = "X") THEN
    thi! = thi! + 1
    IF (thi! > 90) THEN thi! = 90
    REM SET REF3 AT ABOUT thi% FS IN MSB
    temp\% = CINT((whtlv13\% - blklv13\%) / 16 * (thi! / 100))
    POKE &H35, temp%
    REM SET REF6 AT ABOUT thi% FS IN MSB
    temp\% = CINT((whtlv16\% - blklv16\%) / 16 * (thi! / 100))
    POKE &H31, temp%
    POKE &H38, 0: REM UPDATE DAC'S
END IF
IF (key$ = "c") THEN
     flo! = flo! - 1
     IF (flo! < 20) THEN flo! = 20
END IF
```

```
IF (key$ = "C") THEN
     flo! = flo! + 1
     IF (flo! > 70) THEN flo! = 70
END IF
IF (\text{key} = \text{"v"}) THEN
     fhi! = fhi! - .1
     IF (fhi! < .3) THEN fhi! = .3
END IF
IF (key$ = "V") THEN
    fhi! = fhi! + .1
     IF (fhi! > 10) THEN fhi! = 10
END IF
IF (key$ = "b") THEN
     wlo! = wlo! - .1
     IF (wlo! < .5) THEN wlo! = .5
END IF
IF (key$ = "B") THEN
     wlo! = wlo! + .1
     IF (wlo! > 1.5) THEN wlo! = 1.5
END IF
IF (\text{key} = "n") THEN
     whi! = whi! - .1
     IF (whi! < .5) THEN whi! = .5
END IF
IF (key\$ = "N") THEN
     whi! = whi! + .1
     IF (whi! > 1.5) THEN whi! = 1.5
END IF
```

```
IF (key\$ = "m") THEN
    k0! = k0! / 1.1
    IF (k0! < .05) THEN k0! = .05
END IF
                                The Francisco
                                                       10 1 10
IF (key$ = "M") THEN
    k0! = k0! * 1.1
    IF (k0! > .999) THEN k0! = .999
END IF
IF (key\$ = "p") THEN
    dband! = dband! / 1.1
    IF (dband! < .005) THEN dband! = .005
END IF
IF (key$ = "P") THEN
    dband! = dband! * 1.1
    IF (dband! > .35) THEN dband! = .35
END IF
IF (key\$ = "r") THEN
    IF (steps > -6) THEN
          steps = steps - 1
          rg0! = rg0! - .035
          rgl! = rgl! + .0007
          rg2! = rg2! - .0004
    END IF
END IF
IF (key\$ = "R") THEN
     IF (steps < 6) THEN
          steps = steps + 1
          rg0! = rg0! + .035
          rgl! = rgl! - .0007
          rg2! = rg2! + .0004
     END IF
```

END IF

REM UPDATE GAIN CONTROL PARAMETER DISPLAY

COLOR 8

LOCATE 5, 19

PRINT "z(-) Z(+) x(-) X(+) c(-) C(+) v(-) V(+)"

LOCATE 6, 20

format1\$ = "&##&##&##.#"

COLOR 4

PRINT USING formatls; "tlo="; tlo!; " thi="; thi!; " flo = "; flo!; " fni = "; fhi!

1. (4.

 $\mathcal{H}_{\mathcal{F}_{n+1}} = \{\mathcal{H}_{n+1}, \dots, \mathcal{H}_{n+1}, \dots, \mathcal{H}_{n+1}\}$ 

LOCATE 8, 19

COLOR 8

PRINT "b(-) B(+) n(-) N(+) m(-) M(+) p(-) P(+)"

LOCATE 9, 20

format2\$ = "&#.#&#.#&#.###&#.###"

COLOR 4

PRINT USING format2\$; "wlo="; wlo!; " whi="; whi!; " k0="; k0!; " dband="; dband!

**LOCATE 11, 24** 

COLOR 8

PRINT "v-----v'

**LOCATE 12, 23** 

format6\$ = "&+#.####&+#.###&+#.##^^^"

COLOR 4

PRINT USING format6\$; "rg0="; rg0!; " rg1="; rg1!; " rg2="; rg2! . . . . .

RETURN

The embodiments of the invention in which an exclusive property or privilege is claimed are defined as follows:

1. An imaging system for fluorescence endoscopy, comprising: a light source that produces fluorescence excitation light;

an endoscope that delivers the fluorescence excitation light to tissue under examination in vivo and collects autofluorescence produced by the tissue;

- a dual channel fluorescence camera containing a first and second high sensitivity imaging device that receive the autofluorescence in a first and second spectral band and produce electronic signals that are representative of the tissue under examination;
- a control center, including an image processing board that receives the electronic signals produced by the dual channel fluorescence camera and said control center causes an image of the tissue under examination to be processed, stored and displayed on a video monitor;

an automatic gain control circuit that determines a distribution of intensity levels in the electronic signals produced by the dual channel fluorescence camera and adjusts a gain of the first and second high sensitivity image device and/or adjusts a light source intensity based on said distribution of the intensity levels such that the relative gain between the two intensified CCD transducers follows substantially a polynomial; and

- a video monitor that receives the video signals produced by the image processing board and displays an image of the tissue under examination.
- 2. The imaging system for fluorescence endoscopy of Claim 1, wherein the automatic gain control circuit comprises:

a plurality of time-over-threshold counters that determine an image area in one or more video fields that have intensities above a plurality of predetermined thresholds.

- 3. The imaging system for fluorescence endoscopy of Claim 2, wherein the time-over-threshold counters further comprise:
- a clock signal having a frequency substantially equal to a pixel clock of the dual channel fluorescence camera;

- a gating circuit that passes the clock signal during an active portion of the electronic signals produced by the dual channel fluorescence camera;
  - a plurality of counters that count pulses of the gated clock signal;
- a plurality of comparators having the electronic signals produced by the dual channel fluorescence camera connected to a first input and a programmable reference voltage connected to another input such that when the magnitude of the video signals exceed the reference voltage of the comparator, the comparator produces an output which enables one of the plurality of counters; and
- a processor that is programmed to adjust a gain of the high sensitivity imaging device and to adjust the light source intensity such that the distribution of intensity levels in one or more electronic fields substantially equals a desired distribution.
- 4. The imaging system for fluorescence endoscopy of Claim 1, wherein the light source is programmable to produce fluorescence excitation light or white light, the system further comprising:
  - a color video camera coupled to receive light collected by the endoscope;
- a light path directing mechanism selectively positioned to direct light collected by the endoscope to the dual channel fluorescence camera or to the color video camera:
- at least one switch that produces a signal that is indicative of the position of the light path directing mechanism; and
- a light source controller that receives the signal from the switch and causes the light source to produce white light after the signal produced by the switch indicates that the light path directing mechanism is positioned to direct the light collected by the endoscope to the color video camera.
- 5. The imaging system for fluorescence endoscopy of Claim 4, wherein the light source controller causes the light source to stop producing white light and begin producing fluorescence excitation light before the light path directing mechanism is moved from a position where light collected from the endoscope is directed to the color video camera head to a position where light collected from the endoscope is directed to the dual channel fluorescence camera.
- 6. The imaging system for fluorescence endoscopy of Claim 1, wherein the first high sensitivity imaging device receives autofluorescence in a first spectral band and the second high sensitivity imaging device receives autofluorescence in a second spectral band, the imaging system further comprising a central processing unit

that produces a quantitative indication of the intensity of the autofluorescence light in the first spectral band versus the intensity of autofluorescence light in the second spectral band.

7. An imaging system for white light and fluorescence endoscopy, comprising:

a light source that produces white light and fluorescence excitation light;

an endoscope that delivers the light to tissue under examination in vivo and collects reflected light or autofluorescence light produced by the tissue sample;

a fluorescence camera containing a first and second high sensitivity imaging device that receive the autofluorescence in a first and second spectral band and produce electronic signals that are representative of the tissue under examination;

a color video camera that receives the reflected illumination light collected by the endoscope and produces electronic signals that are representative of the tissue under examination;

a control center, including an image processing board, that receives the electronic signals produced by the dual channel fluorescence camera or the color video camera and said control center causes an image of the tissue under examination to be processed, stored and displayed on a video monitor;

a two-part mode switch mechanism including,

- i) a light director that is selectively positioned to direct light collected by the endoscope to the fluorescence camera head or to the color video camera head, and
- ii) a mechanism that operates to change the light source to produce either fluorescence excitation light or white light according to the position of the light director; and

a video monitor that receives the signals produced by the image processing board and displays an image of the tissue under examination.

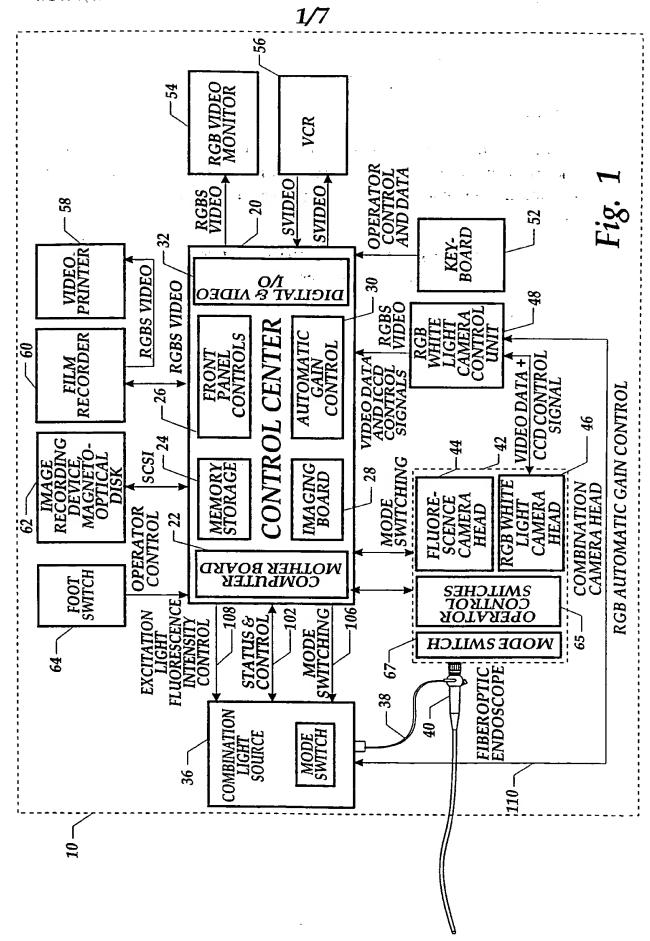
8. The imaging system for white light and fluorescence endoscopy of Claim 7, further comprising:

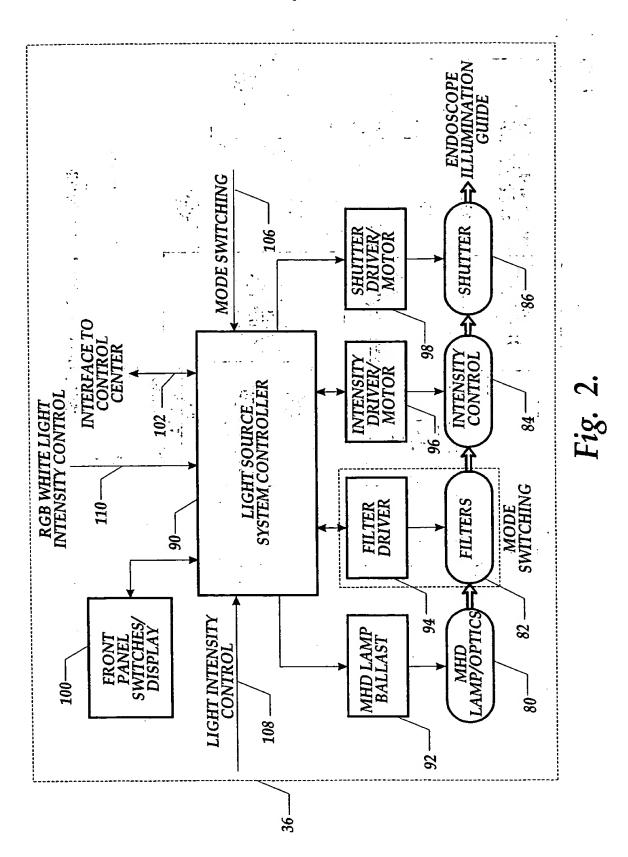
an automatic gain control circuit within the control center that determines a distribution of intensity levels in the electronic signals produced by the first and second high sensitivity imaging devices and adjusts a gain of the high sensitivity imaging devices or adjusts the light source intensity based on said distribution of intensity levels such that the relative gain between the two intensified CCDs follows substantially a polynomial.

- 9. The imaging system for white light and fluorescence endoscopy of Claim 8, wherein the automatic gain control circuit comprises:
- a plurality of time-over-threshold counters that determine an image area in one or more video fields that have intensities above a plurality of predetermined thresholds.
- 10. The imaging system for white light and fluorescence endoscopy of Claim 9, wherein the time-over-threshold counters further comprise:
- a clock signal having a frequency substantially equal to a pixel clock of the intensified CCD transducers;
- a gating circuit that passes the clock signal during an active portion of the electronic signals produced by the dual channel fluorescence camera;
  - a plurality of counters that count pulses of the gated clock signal;
- a plurality of comparators having the electronic signals produced by the dual channel fluorescence camera connected to a first input and a programmable reference voltage connected to another input such that when the magnitude of the electronic signals exceed the reference voltage of the comparator, the comparator produces an output which enables one of the plurality of counters; and
- a processor that is programmed to adjust a gain of the high sensitivity imaging devices and to adjust the light source intensity such that the distribution of intensity levels in one or more video fields substantially equals a desired distribution.
  - 11. An imaging system for fluorescence endoscopy, comprising:
  - a light source that produces fluorescence excitation light;
- an endoscope that delivers the excitation light to tissue under examination in vivo and collects autofluorescence produced by the tissue sample;
- a dual channel fluorescence camera containing a first and second high sensitivity imaging device that receives the autofluorescence light in a first and second spectral band and produces electronic signals that are representative of the tissue sample under examination;
- a control center including an image processing board that receives the electronic signals produced by the dual channel fluorescence camera that are representative of the tissue under examination and produces an indication of a relative intensity of the autofluorescence light in each of the first and second spectral band that produces a portion of an image of the tissue; and

a video monitor that displays the video signals to create an image of the tissue under examination.

- 12. The imaging system for fluorescence endoscopy of Claim 11, wherein the indication of the relative intensity comprises a numeric representation of the relative intensities of the autofluorescence light in each of the first and second spectral bands.
- 13 The imaging system for fluorescence endoscopy of Claim 12, wherein the indication is displayed on a video monitor.





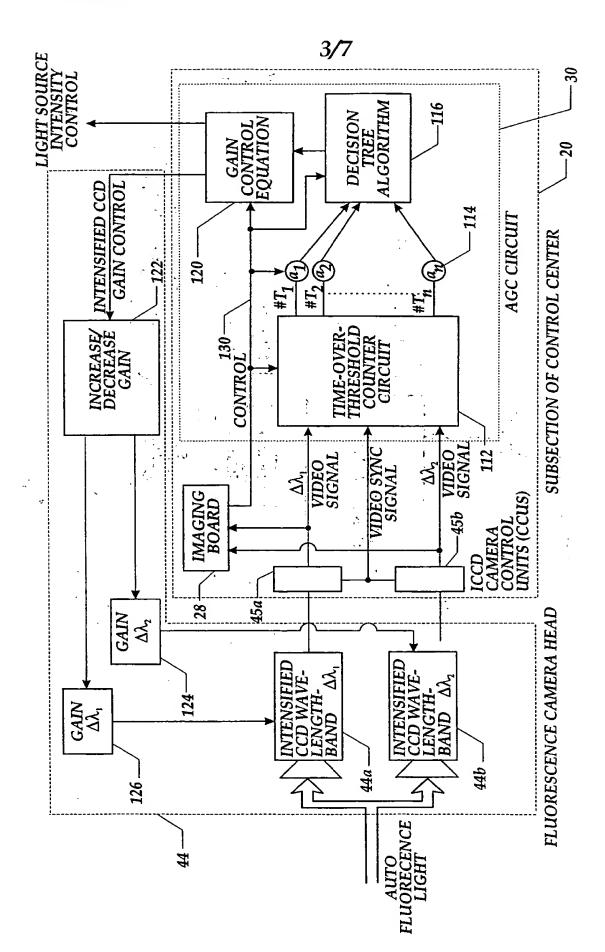
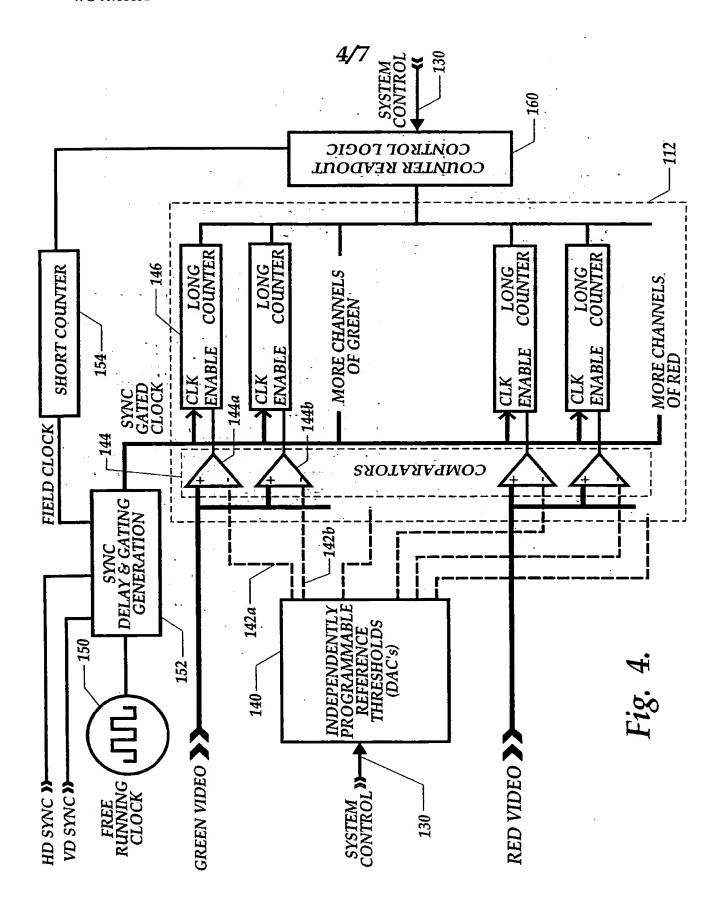
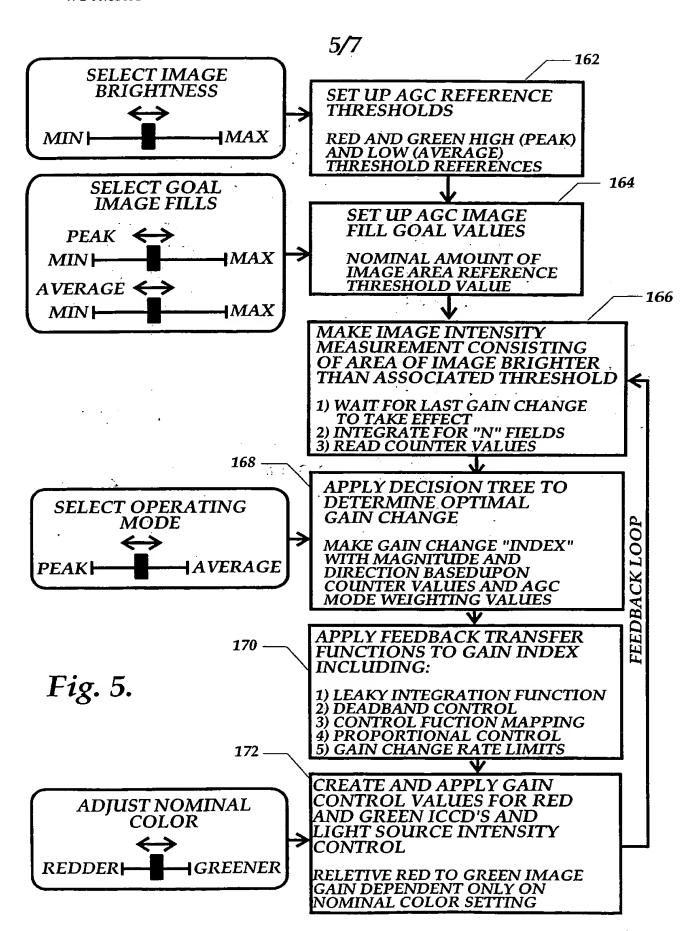


Fig. 3.

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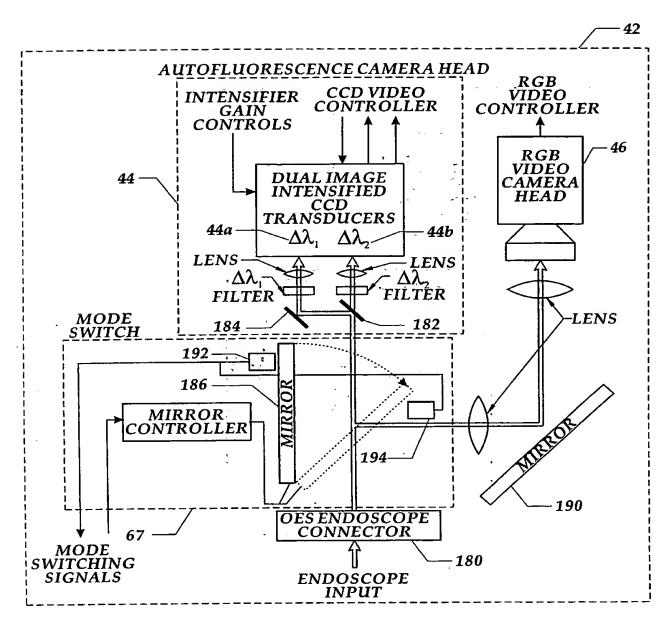
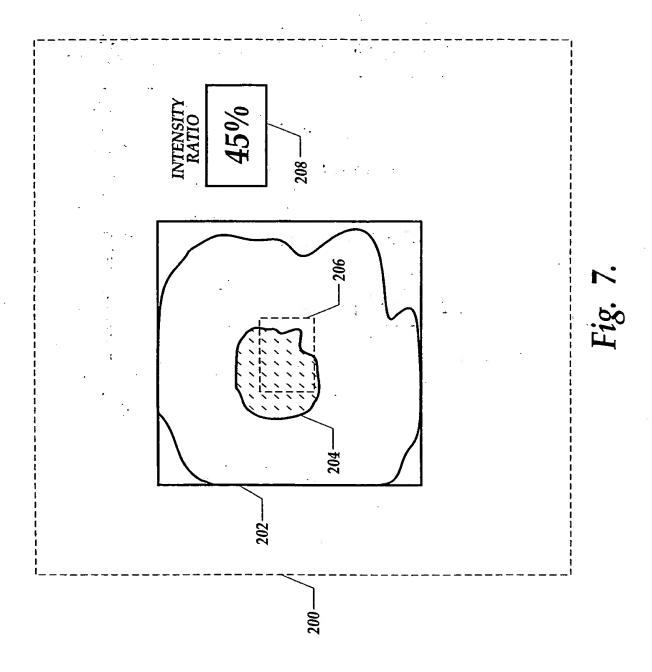


Fig. 6.



## INTERNATIONAL SEARCH REPORT

International Application No PCT/US 99/07789

A. CLASSIF	FICATION OF SUBJECT MATTER		
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C DOCUME	ENTS CONSIDERED TO BE RELEVANT	<u> </u>	
Category 3	Citation of document, with indication, where appropriate, of the	relevant passages	Relevant to claim No.
Υ	WO 95 26673 A (XILLIS)	ţ	1,2,4-9,
	12 October 1995 (1995-10-12)		11-13
	the whole document		
Υ	EP 0 774 865 A (SANYO ELECTRIC)		1,2,4-9,
	21 May 1997 (1997-05-21)		11-13
	column 8, line 12 - line 22 column 21, line 39 - column 22	line 50	
		, 11116 30	
Α	FR 2 671 405 A (INSERM)		7
	10 July 1992 (1992-07-10)		
	page 11, line 10 - page 12, li	ile 3	
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Furt	thar documents are listed in the continuation of box C.	Patent family members are listed	n annex.
* Special ca	ategories of cited documents:	"T" later document published after the inte	mational filing date
"A" docum	tent defining the general state of the art which is not	or priority date and not in conflict with cited to understand the principle or the	eory underlying the
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"P" docum	nent published prior to the international filing date but than the priority date claimed	in the art. "&" document member of the same patent	family
	e actual completion of the international search	Date of mailing of the international sec	
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<u></u>	mailing address of the ISA	Authorized officer	
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	Tel. (+31-70) 340-2040, Tx. 31 651 epo nl, Fax: (+31-70) 340-3016	Lemercier, D	
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## INTERNATIONAL SEARCH REPORT

Information on patent family members

International Application No
PCT/US 99/07789

Patent document cited in search report	t	- Publication date	1	Patent family member(s)	Publication date
WO 9526673	Α	12-10-1995	US	5590660 A	07-01-1997
	•		EP	0752825 A	15-01-1997
			EP	0920831 A	09-06-1999
			JP	10500588 T	20-01-1998
			US	5827190 A	27-10-1998
EP 774865	Α	21-05-1997	JP	.9149308 A	06-06-1997
			JP	10173979 A	26-06-1998
FR 2671405	Α.	10-07-1992	DE	69200916 D	26-01-1995
		•	DE	69200916 T	13÷04-1995
			EP	0565599 A	20-10-1993
			WO	9212412 A	23-07-1992
			JP	6503979 T	12-05-1994
			· US	5370119 A	06-12-1994